

ppln No.: 09/944,326

Amendment Dated: June 17, 2004

Reply to Office Action of February 17, 2004

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1. (currently amended) A <u>pharmaceutical</u> composition for delaying progression of prostatic tumor cells to an androgen- independent state, comprising an antisense oligonucleotide which inhibits expression of TRPM-2 by the tumor cells, whereby when prostatic tumor cells are treated with the composition the progression to androgen independence is delayed, and

a pharmaceutically acceptable carrier suitable for human administration for providing the oligonucleotide to a mammalian subject to reduce expression of TRPM-2.

- 2. The composition of claim 1, wherein the antisense oligonucleotide is complementary to a region of TRPM-2 mRNA including the translation initiation or termination site.
- 3. The composition of claim 1, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 4.
- 4. The composition of claim 1, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 5.
- 5. The composition of claim 1, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 12.
- 6-11 (cancelled)
- 12. (currently amended) The composition of any claim 1, further comprising a second antisense oligodeoxynucleotide which inhibits expression of an anti-apoptotic protein other than TRPM-2.
- 13. The composition of claim 12, wherein the second antisense oligodeoxynucleotide is antisense Bcl-2 oligodeoxynucleotide.
- 14. The composition of claim 12, wherein the antisense oligonucleotide is complementary to a region of TRPM-2 mRNA including the translation initiation or termination site.
- 15. The composition of claim 14, wherein the antisense oligonucleotide is modified to increase the stability of the ODN *in vivo*.

Page 2 of 5

Appln No.: 09/944,326

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- 16. The composition of claim 12, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 4.
- 17. The composition of claim 12, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 5.
- 18. The composition of claim 12, wherein the antisense oligonucleotide has the sequence given by SEQ ID No. 12.
- 19. An oligonucleotide consisting of the sequence set forth in Seq. ID No. 4.
- 20. (canceled)

and 12 and

- 21. An oligonucleotide consisting of the sequence set forth in Seq. ID No. 12.
- 22. (original) A pharmaceutical composition comprising: an antisense oligonucleotide which is complementary to TRPM-2 mRNA and which comprises a continuous sequence of bases as set forth in any of Seq. ID Nos 4, 5

a pharmaceutically acceptable carrier suitable for human administration for providing the olignucleotide to a mammalian subject to reduce expression of TRPM-2.

- 23. (original) The pharmaceutical composition of claim 22, wherein the pharmaceutically acceptable carrier is a lipid carrier.
- 24. (original) The pharmaceutical composition according to claim 22, further comprising an additional antisense oligonucleotide binds specifically to a sequence other than TRPM-2 mRNA.
- 25. (original) The pharmaceutical composition according to claim 24, wherein the additional antisense oligonucleotide binds specifically to a sequence selected from among Bcl-2, Bcl-1x and c-myc.
- 26. (original) The pharmaceutical composition according to claim 25, wherein the additional oligonucleotide consists of the sequence set forth in the Seq. ID No. 13.
- 27. (original) The pharmaceutical composition of claim 22, wherein the antisense oligonucleotide is modified to increase the stability of the ODN *in vivo*.